DATA REVIEW FOR PRIMARY EYE IRRITATION TESTING (870.2400, formerly §81-4)

Product Manager: 23 **Reviewer:** Byron T. Backus, Ph.D.
MRID No.: 45086101 & 45255601

Citation: Tay, C.H. Second Amended Report Date: October 24, 2000, Acute Eye Irritation - OPPTS; Toxicon Corporation, 15 Wiggins Avenue, Bedford, MA 01730; Laboratory Project ID/Study Number 00-0661-G1.

Sponsor: Compliance Services International, Inc., Representative to Becker Underwood, Inc., 1112 Alexander Avenue, Tacoma, WA 98421

Quality Assurance (40 CFR §160.12): Included (p. 6)

Test Material: Admiral WSP; Lot/Batch #: 10673; a dark blue powder; this study was apparently conducted on the formulation as originally proposed for registration (Acid Blue 9 (Erioglaucine): 68.13%; Acid Yellow 23 (Tartrazine): 4.51%).

Dosage: 0.1 mL
Species: Rabbits; New Zealand White
Age: Adult (at least 10 weeks old)
Weight: 2.21-2.27 kg
Source: Millbrook Breeding Labs, Amherst, MA

Executive Summary: In an amended eye irritation study (amended report in MRID 45255601; original report in MRID 45086101) 0.1 mL of the test article was instilled into the left eye of each of 3 New Zealand White rabbits, with subsequent 72-hour observation.

Admiral WSP; Lot/Batch #: 10673 (with active ingredients Acid Blue 9: 68.13% and Acid Yellow 23: 4.51%); is in toxicity category IV for the tested fromulation, based on the lack of irritation (all scores zero) in any of the 3 eyes at 1, 24, 48 and 72 hours.

Although this eye irritation study is classified as acceptable (toxicity category IV) for the tested formulation, because of uncertainties relating to the irritation potential of the revised formulation (active ingredients: Acid Blue 9 [Erioglaucine] at 49.62%, and Acid Yellow 23 [Tartrazine] at 3.05%), TRB recommends that this study be accepted as defining a toxicity category III (not IV) for the revised formulation.

Procedure (including deviations from 870.2400): "Three animals were treated by instilling 0.1 mL of the test article in the left eye of each animal. The right eye was left untreated to serve as control."

"The eyes of the test animals were not washed out for 24 hours following installation of the test substance. After the 24 hour reading, all eyes were washed with 0.9% USP Sodium Chloride for Injection (NaCl)."

Summary:

Observations	Number "positive"/number tested			
	Hours			
	1	24	48	72
Corneal Opacity	0/3	0/3	0/3	0/3
Iritis	0/3	0/3	0/3	0/3
Conjunctivae:				
Redness[a]	0/3	0/3	0/3	0/3
Chemosis[a]	0/3	0/3	0/3	0/3

[a]Score of 2 or more considered to be "positive."
Data summarized from Tables II-A, II-B & II-C, pp. 16-18 of MRID 45255601.

No irritation was observed. In the original report (MRID 45086101) it was stated (p. 13) that: "Slight irritation was observed in all the treated eyes 1 hour after dosing. This irritation was resolved in all the treated eyes by the 48 hour observation point." In this amended report (MRID 45255601) it is stated (p. 19) that: "The sentences 'slight irritation was observed in all the treated eyes 1 hour after dosing> This irritation was resolved in all the treated eyes by the 48 hour observation point.' were removed... These sentences were included in the original report due to a template error and have been removed. The remaining sentence was corrected to accurately reflect the raw data in that no irritation was observed in any of the treated or control eyes at any of the observation points."

ACUTE TOX ONE-LINERS

1. **DP BARCODE:** D271422
2. **PC CODES:** 110301 & 110302; Erioglaucine & Tartrazine
3. **CURRENT DATE:** March 26, 2001
4. **TEST MATERIAL:** Admiral WSP, EPA File Symbol 67064-R. The acute oral LD_{50} study below was conducted on a revised formulation, with active ingredients Acid Blue 9 [Erioglaucine] 49.72% & Acid Yellow 23 [Tartrazine] 3.27%, with other ingredients 47.01%. The eye irritation study below was conducted on the formulation as originally proposed for registration, with active ingredients Acid Blue 9 [Erioglaucine] 68.13% & Acid Yellow 23 [Tartrazine] 4.51%, with other ingredients 27.36%.

Study/Species/Lab Study #/Date	MRID	Results	Tox. Cat.	Core Grade
Acute oral toxicity/rat/MB Research Laboratories/MB Research Project # 00-8731.01/Dec. 5 2000	45281101	LD_{50} > 5000 mg/kg for Admiral WSP (males,females, combined). No mortalities. Symptoms: Emaciation in one animal on days 3-4, blue staining of the anogenital and/or other body regions.	IV	A
Primary eye irritation/rabbit/Toxicon Corporation/Study No. 00-0661-G1/October 24, 2000 (amended report date)	45255601; originally submitted as 45086101	No eye irritation. All scores zero. However, formulation tested has been revised by registrant.	III[a]	A

[a]Registrant has revised the proposed formulation; because of uncertainties involving possible changes in its eye irritation potential, TRB recommends that this study be accepted as defining a toxicity category III (not IV) for the revised formulation.

Core Grade Key: **A =Acceptable, S = Supplementary, U = Unacceptable, V = Self Validated**